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# Why Vaccines for Staph Infections Always Fail

*Staphylococcus aureus bacteria use an immune system trick to colonize up to 80% of humans, and that same strategy to evade vaccine-based immunity. This explains why all staph vaccines up to now have failed to provide clinically relevant immunity, according to a paper published last month in Cell Reports Medicine.*

By Angelo DePalma, Ph.D.

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Research into vaccines for Staphylococcus aureus (*S. aureus*), the most common type of staph infection, has led to experimental vaccines that — like flu vaccines — fail in humans. A paper published Jan. 16 in Cell Reports Medicine explained why.

When a person first encounters staph, the bacterium fools the human immune system into releasing ineffective antibodies instead of the neutralizing antibodies typically associated with robust immunity. That "trick" allows *S. aureus* to colonize us, usually harmlessly.

When a colonized person's immune system is later challenged with a staph vaccine it does not make new, effective antibodies. Instead, it calls up more of the same ineffective antibodies that allowed the bug to colonize the individual in the first place.

Vaccine developers have tried at least three different approaches to *S. aureus* vaccination but all were met with the same issue.

**The immune system is willing ...**

*S. aureus* is one of 30 Staphylococcus species in nature and 11 that colonize humans as part of the human microbiome. It is found in the nostrils, skin and other reservoirs in healthy people and is only dangerous when it enters the bloodstream, particularly in immunocompromised individuals.

Up to 80% of humans harbor Staphylococcus species.

The human immune system makes antibodies against *S. aureus* as it does against other microbial invaders. But instead of neutralizing antibodies that fight colonization and infection, it produces antibodies that allow the bug to continue colonizing us.

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When the colonized person is challenged with either *S. aureus* infection or vaccination, these dummy antibodies interfere in force but do nothing to help the patient. Vaccination amplifies this effect — which is why *S. aureus* infections must be treated with antibiotics.

Vaccination "only works when the initial immune response to that pathogen was actually protective," said J.R. Caldera, Ph.D., a co-author of the paper, in a news release.

Some 80% of staph infections are caused by the invasion of the same strain colonizing the individual's nose or skin, their "initial immune response" was not protective so subsequent responses will not be either.

"What sets SA (Staphylococcus aureus) apart is that the bacteria themselves have ways of enabling the immune system from the moment they encounter us, and these evasive strategies are only reinforced by vaccination," Caldera said.

**... but the antibodies are weak**

Anti-staph vaccines generate strong immune responses to vaccination and infection but those responses are directed toward bacterial features that *S. aureus* uses to fool its host into accepting peaceful coexistence.

This is a case of vaccine-induced immune system priming or "original antigenic sin" — the process by which the bacterium locks in the unsuccessful people make when confronted with the pathogen.

This failure eventually led vaccine researchers down another dark alley, toward vaccines targeting the *S. aureus* toxins listed in the bacterium's arsenal. So-called "toxoid" strategies are the basis of tetanus, diphtheria and DTP vaccines.

But "remarkably, both active (vaccine-based) and passive (antibody-based) platforms of immunization against SA toxins were met with failures," said senior author Georgia Liu M.D., Ph.D., professor of pediatrics at the University of California San Diego School of Medicine.

For example, a 2021 AstraZeneca-funded study of suvratoxumab, a monoclonal antibody targeting the *S. aureus* toxin, found that progression to pneumonia in staph-infected patients was no lower in treated than in untreated subjects.

Most pathogens, especially bacteria, are complex organisms carrying several different antigens. Vaccine developers usually target the most prominent antigen to trigger the strongest, most relevant immune response.

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On that basis, Liu considered a third possible *S. aureus* vaccine strategy: targeting minor cell wall antigens on *S. aureus* instead of its toxins or the main antigen.

This approach would tend to induce weaker immune responses requiring high vaccine or adjuvant doses, but it fell short as well.

**Antibiotic resistance leads to 'super-bugs'**

Nostrils are the main staph reservoir in humans and a significant source of infection, but the bug also colonizes the skin, perineum, vagina, throat and gastrointestinal tract.

Staph infections occur when the bacteria enter the bloodstream, joints, heart or skin, usually when the person's immune system is weakened. Standard antibiotics usually cure staph infections.

However, over the past 70 years, bacteria that colonize humans have found ways to counter the use or overuse of antibiotics and antimicrobials designed to kill them. Some bacteria have developed resistance to these agents, making antibiotics less effective or completely ineffective.

One type of antibiotic-resistant *S. aureus*, "methicillin-resistant" *S. aureus* or "MRSA," is of particular concern.

The most common MRSA outcome outside of hospitals is a skin infection. But serious cases can lead to pneumonia or other serious organ infections. Untreated MRSA infections can cause sepsis — an extreme, system-wide response to an infection.

Hospitalized patients are more susceptible to severe, life-threatening outcomes since they are exposed to fellow patients' staph strains as well as the ones they carry. Surgical site infections are a significant source of serious, systemic staph infections.

The medical and social costs, direct and indirect, of antibiotic resistance in the U.S. may be as high as $55 billion per year. More than 2.8 million Americans per year have an antibiotic-resistant infection and 35,000 die. *S. aureus* caused nearly 120,000 bloodstream infections — the most serious kind — and 20,000 deaths in the U.S. in 2017.

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**Could *S. aureus* be beneficial?**

The negative effects of *S. aureus* on human health are fairly well understood.

We know staph bacteria colonize us, are tolerated by the immune system and cause disease when they enter the bloodstream or other sites in the body. We know that *S. aureus* antibody responses do not clear the bacterium or eliminate either colonization or infection.

But the role of *S. aureus* as part of a normal, healthy microbiome has not been extensively investigated.

A 2022 study on components of the skin microbiome suggested that at least one Staphylococcus species, *S. hominis* — the bug mostly responsible for body odor — may prevent skin infections.

Another species that mainly colonizes skin, *S. epidermis*, is both anti-inflammatory and antibacterial.

A 2015 study found that chronic *S. aureus* infection prevented the development of autoimmune encephalomyelitis in a rat model of multiple sclerosis. Encephalomyelitis is inflammation of the brain and spinal cord. Although infection itself caused some types of inflammatory markers to rise it reduced the severity of nerve cell and central nervous system inflammation.

"SA [*S. aureus*] has been with humans a long time, so it's learned how to be part-time symbiont, part-time deadly pathogen," Liu said. "If we're going to develop effective vaccines against SA, we need to understand and overcome the strategies it uses to maintain this lifestyle."

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Angelo DePalma, Ph.D.
Angelo DePalma, Ph.D., is a science reporter/editor for The Defender.

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Plenty of "meat" when you read about it. Although there is no strong evidence that echinacea has health benefits in any area, echinacea is most commonly used to treat colds. Some early research shows that echinacea supplements may have broader health benefits as well.
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4 days ago
Yes, Echinacea has been proven to shorten the duration of colds and respiratory infections if taken early.
Echinacea has also been shown to help with skin infections and help keep skin looking clear ... I have experienced this over the years first-hand, although I do not take it on a regular basis.
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I have heard of many tragic cases of walking, talking, normal children who wound up with profound mental disorders after vaccines.' Rand Paul
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No vaccine is worthy of use. Your body always knows best if you allow it to work as it wants to. Big pharma drugs will always disrupt and poison that process.
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And yet another 'vaccine' needed???? Oh puleeez.
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The Arch Enmity of Jabbers strikes again. Those peanuts misbehave all the time when cut with Tadd Pharma says "It's not our product it was Victim!" Our collective expert scientific brains are no match for Devious More-Intelligent Bacteria!
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4 days ago
Interesting. Another proof that mice are not humans when it comes to medical experiments of safety and efficacy.
Anyone with a weakened immune system, typically if they have underlying heart or lung disease," is at risk ... comorbi CTE jab.
Yet another case of vaccine-induced similar bacteria.
"Cases of invasive group A strep rising across the country"
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Here is some interesting information:
In 2015, we reported the antimicrobial treatment of *S. aureus* infections, in the present study, we screened three known antibacterials (CPR, ALB, and LEV) and found that CPR but not ALB or LEV had a potent anti-bacterial activity against *S. aureus* species. Our novel findings indicate that CPR treatment has an anti-bacterial effect against certain *S. aureus* isolates. However, to comprehend why ivermectin did not inhibit the growth of all Staphylococci needs further investigation. Nevertheless, we have extended the broad-range of known pharmacological effects of ivermectin. As pharmacological screening of ivermectin are being carried out in clinical development as an anti-staphylococcal agent is potentially ongoing.
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